

## AMENDMENTS

### **A Listing of the Claims**

The following listing of claims replaces all previous listings or versions thereof:

1-63 (Cancelled)

64. (Previously presented) A method of comparing one or more nucleic acid targets within two or more samples, comprising:

- a) preparing a sample mixture by a process comprising obtaining at least a first sample and a second sample, each potentially having at least a first nucleic acid target and mixing the first nucleic acid sample and the second nucleic acid sample to create a sample mixture;
- b) isolating at least a first target fraction of the sample mixture;
- c) performing at least a first amplification reaction on the first target fraction, wherein the amplification reaction produces at least a first amplified nucleic acid, if the first nucleic acid target is present in the first sample, and at least a second amplified nucleic acid, if the first nucleic acid target is present in the second sample;
- d) differentiating the first amplified nucleic acid in the first target fraction, if any, from the second amplified nucleic acid in the first target fraction, if any; and
- e) comparing abundance of the first nucleic acid target of said first sample to abundance of the first nucleic acid target of said second sample.

65. (Previously presented) The method of claim 64, further defined as comprising performing at least a first amplification reaction on the first target fraction using at least a first target-specific primer, wherein the amplification reaction produces at least a first amplified

nucleic acid, if the first nucleic acid target is present in the first sample, and at least a second amplified nucleic acid, if the first nucleic acid target is present in the second sample.

66. (Previously presented) The method of claim 65, further defined as comprising isolating at least a second target fraction from the sample mixture and performing a second amplification reaction on the the second target fraction using a second target-specific primer that is specific for a second target that may be present in the first sample and/or the second sample.

67. (Previously presented) The method of claim 64, wherein:

a) preparing the sample mixture is further defined as comprising:

preparing at least a first tagged nucleic acid sample by appending at least a first nucleic acid tag comprising a first amplification domain and a first differentiation domain to the first nucleic acid target of the first sample, if the first nucleic acid target is present in the first sample;

preparing at least a second tagged nucleic acid sample by appending at least a second nucleic acid tag comprising a second amplification domain and a second differentiation domain to the first nucleic acid target of the second sample, if the first nucleic acid target is present in the second sample, wherein the second differentiation domain is different from the first differentiation domain; and

mixing the first tagged nucleic acid sample and the second tagged nucleic acid sample to create the sample mixture; and

b) performing at least a first amplification reaction on the first target fraction is further defined as producing at least a first amplified nucleic acid comprising the first differentiation domain and a segment of the first nucleic acid target, if the first nucleic acid target is present in the first sample, and at least a second

amplified nucleic acid comprising the second differentiation domain and a segment of the first nucleic acid target, if the first nucleic acid target is present in the second sample.

68. (Previously presented) The method of claim 67, wherein the differentiation domain of the first tag and the second tag is appended between the first nucleic acid target sequence and the amplification domain.

69. (Previously presented) The method of claim 67, wherein said nucleic acid target is one target of a plurality of nucleic acid targets within the samples.

70. (Previously presented) The method of claim 67, wherein said first and second samples are two samples of a plurality of samples.

71. (Previously presented) The method of claim 70, wherein the first and second tag are two tags of a plurality of tags.

72. (Previously presented) The method of claim 64, wherein the amplification domains of the first and second nucleic acid tags are functionally equivalent.

73. (Previously presented) The method of claim 72, wherein the amplification domains of the first and second nucleic acid tags are identical.

74. (Previously presented) The method of claim 67, wherein the amplification domain of the first nucleic acid tag and the second nucleic acid tag comprises a primer binding domain.

75. (Previously presented) The method of claim 67, wherein the amplification domain of the first nucleic acid tag and the second nucleic acid tag comprises a transcription domain.

76. (Previously presented) The method of claim 67, wherein the differentiation domain of the first nucleic acid tag and the second nucleic acid tag comprise at least a size differentiation domain, an affinity domain, or a unique sequence domain.
77. (Previously presented) The method of claim 67, wherein said first target fraction is one of a plurality of target fractions.
78. (Previously presented) The method of claim 67, wherein the first target fraction is isolated by binding a ligand to at least a segment of the first nucleic acid target.
79. (Previously presented) The method of claim 78, wherein the ligand is a nucleic acid, protein, or other molecule with an affinity for certain nucleic acids.
80. (Previously presented) The method of claim 79, wherein the ligand is a nucleic acid complementary to at least a segment of the first nucleic acid target.
81. (Previously presented) The method of claim 80, wherein the first complementary nucleic acid is used to separate the first target nucleic acid from at least one other nucleic acid or molecule.
82. (Previously presented) The method of claim 81, wherein the target fraction is subsequently removed from the first complementary nucleic acid.
83. (Previously presented) The method of claim 80, wherein the first complementary nucleic acid is one of a plurality of complementary nucleic acids, and each complementary nucleic acid is complementary to one of a plurality of nucleic acid targets.
84. (Previously presented) The method of claim 80, wherein the first complementary nucleic acid is bound to a solid support.

85. (Previously presented) The method of claim 84, wherein the first complementary nucleic acid is one of a plurality of complementary acids bound to an array, and each of the complementary nucleic acids is complementary to one of a plurality of nucleic acid targets.

86. (Previously presented) The method of claim 84, wherein the solid support is one of a plurality of solid supports.

87. (Previously presented) The method of claim 84, wherein the solid support is an array, a microtiter well, a chip, a bead or a combination thereof.

88. (Previously presented) The method of claim 67, wherein differentiating comprises binding the first amplified nucleic acid to a ligand specific to at least a segment of the first differentiation domain or binding the second amplified nucleic acid to at least a segment of the second differentiation domain.

89. (Previously presented) The method of claim 67, wherein said differentiation domain of the first nucleic acid tag comprises a first affinity domain and the second nucleic acid tag comprises a second affinity domain that is distinct from the first affinity domain.

90. (Previously presented) The method of claim 89, wherein differentiating comprises binding at least a segment of the first affinity domain to a first affinity domain specific ligand and/or binding at least a segment of the second affinity domain to a second affinity domain specific ligand.

91. (Previously presented) The method of claim 88, wherein the first and second affinity domain specific ligands are two of a plurality of ligands.

92. (Previously presented) The method of claim 90, wherein at least one of the first or the second affinity domain specific ligands is labeled.

93. (Previously presented) The method of claim 90, wherein the binding of the first affinity domain specific ligand to the first affinity domain results in a detectable signal and/or the binding of the second affinity domain specific ligand to the second affinity domain results in a detectable signal.

94. (Previously presented) The method of claim 93, wherein the binding of the first affinity domain specific ligand to the first affinity domain results in a first detectable signal and the binding of the second affinity domain specific ligand to the second affinity domain results in a second detectable signal.

95. (Previously presented) The method of claim 94, wherein the first detectable signal is distinguishable from the second detectable signal.

96. (Previously presented) The method of claim 67, wherein the differentiation domain of the first nucleic acid tag comprises a first sequence domain and the differentiation domain of the second nucleic acid tag comprises a second sequence domain that is distinguishable from the first sequence domain.

97. (Previously presented) The method of claim 67, wherein differentiating comprises sequencing the first amplified nucleic acid and the second amplified nucleic acid.

98. (Previously presented) The method of claim 67, wherein the differentiation domain is a unique size domain.

99. (Previously presented) The method of claim 67, wherein the first nucleic acid tag or the second nucleic acid tag further comprises at least one additional domain.

100. (Previously presented) The method of claim 99, wherein said additional domain is a restriction enzyme domain, a secondary amplification domain, a sequencing primer binding site, a labeling domain or a combination thereof.

101. (Previously presented) The method of claim 99, wherein said additional domain comprises one or more restriction enzyme domains.

102. (Previously presented) The method of claim 99, wherein said additional domain comprises a labeling domain.

103. (Previously presented) The method of claim 102, wherein the labeling domain is a transcription promoter.

104. (Previously presented) The method of claim 102, wherein the labeling domain is a primer binding site.

105. (Previously presented) A method of comparing one or more nucleic acid targets within two or more samples, comprising:

- a) obtaining at least a first sample and a second sample, each potentially having at least a first nucleic acid target;
- b) preparing at least a first tagged nucleic acid sample by appending at least a first nucleic acid tag comprising a first amplification domain and a first differentiation domain to the first nucleic acid target of the first sample, if the first nucleic acid target is present in the first sample;
- c) preparing at least a second tagged nucleic acid sample by appending at least a second nucleic acid tag comprising a second amplification domain and a second differentiation domain to the first nucleic acid target of the second sample, if the first nucleic acid target is present in the second sample;
- d) mixing the first tagged nucleic acid sample and the second tagged nucleic acid sample to create a sample mixture;

- e) isolating at least a first target fraction of the sample mixture;
- f) performing at least a first amplification reaction on the first target fraction, wherein the amplification reaction produces at least a first amplified nucleic acid comprising the first differentiation domain and a segment of the first nucleic acid target, if the first nucleic acid target is present in the first sample, and at least a second amplified nucleic acid comprising the second differentiation domain and a segment of the first nucleic acid target, if the first nucleic acid target is present in the second sample;
- g) differentiating the first amplified nucleic acid in the first target fraction, if any, from the second amplified nucleic acid in the first target fraction, if any; and
- h) comparing the first nucleic acid target of said first sample to the nucleic acid target of said second sample.

106. (Previously presented) A method of comparing one or more nucleic acid targets within two or more samples comprising

- a) obtaining at least a first sample and a second sample, each potentially having at least a first nucleic acid target;
- b) preparing at least a first tagged nucleic acid sample by appending at least a first nucleic acid tag comprising a first amplification domain and a first differentiation domain to the first nucleic acid target of the first sample, if the first nucleic acid target is present in the first sample, wherein the first differentiation domain comprises a first affinity domain;
- c) preparing at least a second tagged nucleic acid sample by appending at least a second nucleic acid tag comprising a second amplification domain and a second differentiation domain to the first nucleic acid target of the second sample, if the



first nucleic acid target is present in the second sample, wherein the first differentiation domain comprises a second affinity domain that is distinct from the first differentiation domain;

- d) mixing the first tagged nucleic acid sample and the second tagged nucleic acid sample to create a sample mixture;
- e) isolating at least a first target fraction of the sample mixture;
- f) performing at least a first amplification reaction on the first target fraction, wherein the amplification reaction produces at least a first amplified nucleic acid comprising the first affinity domain and a segment of the first nucleic acid target, if the first nucleic acid target is present in the first sample, and at least a second amplified nucleic acid comprising the second affinity domain and a segment of the first nucleic acid target, if the first nucleic acid target is present in the second sample;
- g) differentiating the first amplified nucleic acid in the first target fraction, if any, from the second amplified nucleic acid in the first target fraction, if any, by binding the first affinity domain to a first ligand and the second affinity domain to a second ligand; and
- h) comparing the first nucleic acid target of said first sample to the nucleic acid target of said second sample.